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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/989,728	11/20/2001	Avi J. Ashkenazi	P2730P1C72	2424
35489	7590	10/19/2004	EXAMINER	
HELLER EHRMAN WHITE & MCAULIFFE LLP 275 MIDDLEFIELD ROAD MENLO PARK, CA 94025-3506			HAMUD, FOZIA M	
			ART UNIT	PAPER NUMBER
			1647	

DATE MAILED: 10/19/2004

Please find below and/or attached an Office communication concerning this application or proceeding.

Office Action Summary

Application No.	Applicant(s)	
09/989,728	ASHKENAZI ET AL.	
Examiner	Art Unit	
Fozia M Hamud	1647	

-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --

Period for Reply

A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) FROM THE MAILING DATE OF THIS COMMUNICATION.

- Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication.
- If the period for reply specified above is less than thirty (30) days, a reply within the statutory minimum of thirty (30) days will be considered timely.
- If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication.
- Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).

Status

1) Responsive to communication(s) filed on 25 August 2003.
2a) This action is **FINAL**. 2b) This action is non-final.
3) Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under *Ex parte Quayle*, 1935 C.D. 11, 453 O.G. 213.

Disposition of Claims

4) Claim(s) 119-138 is/are pending in the application.
4a) Of the above claim(s) _____ is/are withdrawn from consideration.
5) Claim(s) _____ is/are allowed.
6) Claim(s) 119-138 is/are rejected.
7) Claim(s) _____ is/are objected to.
8) Claim(s) _____ are subject to restriction and/or election requirement.

Application Papers

9) The specification is objected to by the Examiner.
10) The drawing(s) filed on 21 November 2001 is/are: a) accepted or b) objected to by the Examiner.
 Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).
 Replacement drawing sheet(s) including the correction is required if the drawing(s) is objected to. See 37 CFR 1.121(d).
11) The oath or declaration is objected to by the Examiner. Note the attached Office Action or form PTO-152.

Priority under 35 U.S.C. § 119

12) Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).
a) All b) Some * c) None of:
 1. Certified copies of the priority documents have been received.
 2. Certified copies of the priority documents have been received in Application No. _____.
 3. Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).

* See the attached detailed Office action for a list of the certified copies not received.

Attachment(s)

1) Notice of References Cited (PTO-892)
2) Notice of Draftsperson's Patent Drawing Review (PTO-948)
3) Information Disclosure Statement(s) (PTO-1449 or PTO/SB/08)
 Paper No(s)/Mail Date _____.
4) Interview Summary (PTO-413)
 Paper No(s)/Mail Date _____.
5) Notice of Informal Patent Application (PTO-152)
6) Other: _____.

DETAILED ACTION

1. Applicant's preliminary amendment canceling claims 1-118 and adding new claims 119-138, filed on 20 November 2001 is acknowledged.

Thus claims 119-138 are pending and under consideration.

2. **Priority:**

2a. Based on the information given by Applicants and an inspection of the patent applications, the Examiner has concluded that the subject matter defined in this application is supported by the disclosure in application serial no. 09/941,992 filed on 28 August 2001, because, EXAMPLE 160 (Assay #111; Chondrocyte proliferation assay which demonstrates that the polypeptide encoded by the claimed nucleic acid, induces the proliferation of chondrocytes), which provides a specific and substantial asserted utility or a well established utility for the claimed nucleic acid is disclosed on page 531 of Application no. 09/941,992. However, it does not appear that any of the other prior applications disclose this assay. Specifically, it does not appear that PCT/US99/12252 filed on 02 June 1999, in which the current application claims priority to, discloses the Chondrocyte proliferation assay. Accordingly, the subject matter defined in claims 119-138, is afforded an effective filing date of 28 August 2001, which is the filing date of the U.S application No. 09/941,992.

Should the applicant disagree with the examiner's factual determination above, it is incumbent upon the applicant to provide the serial number and specific page number(s) of any parent application filed prior to 09/04/01, which specifically supports the particular claim limitation for each and every claim limitation in all the pending claims

which applicant considers to have been in possession of and fully enabled for prior to 08/28/01.

Specification:

3a. The lengthy specification has not been checked to the extent necessary to determine the presence of all possible minor errors. Applicant's cooperation is requested in correcting any errors of which applicant may become aware in the specification.

Information Disclosure Statement:

4a. References A1 and A2, cited on the PTO-1449 form submitted by Applicants on 31 May 2002 have not been considered, because they do not comply with 37 CFR 1.98(a)(2) requirements, since they fail to identify each publication by author and publication date. Applicant is advised that the date of submission of any item of information or any missing element(s) will be the date of submission for purposes of determining compliance with the requirements based on the time of filing the IDS, including all "statement" requirements of 37 CFR 1.97(e). See MPEP § 609 C(1).

Claim rejections-35 USC § 112, first paragraph:

The following is a quotation of the first paragraph of 35 U.S.C. 112:

The specification shall contain a written description of the invention, and of the manner and process of making and using it, in such full, clear, concise, and exact terms as to enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the same and shall set forth the best mode contemplated by the inventor of carrying out his invention.

5a. Claims 119-123 and 132-134 are rejected under 35 U.S.C. 112, first paragraph, while being enabling for an isolated nucleic acid comprising the nucleotide sequence set forth in SEQ ID NO:421, and encoding the polypeptide of SEQ ID NO:422, said

polypeptide which induces the proliferation of chondrocytes, does not reasonably provide enablement for an isolated nucleic acid having at least 80%, 85%, 90%, 95% or 99% identity to the nucleic acid encoding the polypeptide of SEQ ID NO:422. The specification does not enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make or use the invention.

The instant claims 119-123, 132-134 are drawn to an isolated nucleic acid that has at least "80%, 85%, 90%, 95% or 99%" identity to the nucleic acid of SEQ ID NO:421, or having at least 80%, 85%, 90%, 95% or 99% to a nucleic acid encoding the polypeptide of SEQ ID NO:422, or all of the nucleic acids that hybridize to a nucleic acid encoding the polypeptide of SEQ ID NO:422, however, instant specification does not teach how to make or use said nucleic acid. Instant specification discloses that the polypeptide of SEQ ID NO:422 encoded by the claimed nucleic acid induces proliferation of chondrocytes, therefore, said polypeptide is expected to be useful for the treatment of various bone and/or cartilage disorders such as, for example, sports injuries and arthritis, (see Example 160, assay 111 on page 531). Therefore, only the full length polypeptide of SEQ ID NO:422 encoded nucleic acid of SEQ ID NO:421 can be used for said treatments, because Applicants have not shown that variants of the polypeptide of SEQ ID NO:422, induce chondrocyte proliferation.

Instant claims 119-123, 132-134 are drawn to a genus of nucleic acids that are defined only by sequence identity. Due to the large quantity of experimentation necessary to determine all the nucleic acids comprising a nucleotide sequence that is at least 80%, 85%, 90%, 95% or 99% identical to the nucleic acid of SEQ ID NO:421, or

those that hybridize to the nucleic acid of SEQ ID NO:421, and to screen for the ones that encode the polypeptide of SEQ ID NO:422, the lack of direction/guidance presented in the specification regarding which variants of the nucleic acid of SEQ ID NO:421 would retain the desired activity, the complex nature of the invention, the absence of working examples directed to variants of the nucleic acid of SEQ ID NO:421, the state of the prior art establishing that biological activity cannot be predicted based on structural similarity, the unpredictability of the effects of mutation on the structure and function of the claimed polypeptide, and the breadth of the claims which fail to recite particular biological activities, undue experimentation would be required of the skilled artisan to make and/or use the claimed invention in its full scope.

5b. Claims 119-123, 132-134 are rejected under 35 U.S.C. §112, first paragraph, as failing to comply with the written description requirement. The claims contains subject matter which was not described in the specification in such a way as to reasonably convey to one skilled in the relevant art that the inventors, at the time the application was filed, had possession of the claimed invention.

The instant claims 119-123 are drawn to an isolated nucleic acid that shares “80%, 85%, 90%, 95% or 99%” identity to the nucleic acid of SEQ ID NO:421 or to a nucleic acid that encodes the polypeptide of SEQ ID NO:422, and claims 132-134 are drawn to an isolated nucleic acid which hybridize to a nucleic acid encoding a specific polypeptide. However, the instant specification only describes the structure of the nucleic acid of SEQ ID NO:421, and therefore, conception is not achieved until reduction to practice has occurred. Adequate written description requires more than a

mere statement that it is part of the invention. To provide adequate written description and evidence of possession of a claimed genus, the specification must provide sufficient distinguishing identifying characteristics of the genus. The factors to be considered include disclosure of compete or partial structure, physical and/or chemical properties, functional characteristics, structure/function correlation, methods of making the claimed product, or any combination thereof. In this case, the only factor present in the claim is a partial structure in the form of a recitation of percent identity or hybridizing language. There is not even identification of any particular portion of the structure that must be conserved. Accordingly, in the absence of sufficient recitation of distinguishing identifying characteristics, the specification does not provide adequate written description of the claimed genus. Vas-cath Inc. v. Mahurkar, 19 USPQ2d I 1111, clearly states "applicant must convey with reasonable clarity to those skilled in the art that, as of the filing date sought, he or she was in possession of the invention. The invention is, for purposes of the 'written description' inquiry, whatever is now claimed." (See page 1117.) The specification does not "clearly allow persons of ordinary skill in the art to recognize that he or she invented what is claimed." (See Vas-cath at page 1116). As discussed above, the skilled artisan cannot envision the detailed chemical structure of the encompassed genus of polypeptides, and therefore conception is not achieved until reduction to practice has occurred, regardless of the complexity or simplicity of the method of isolation. Adequate written description requires more than a mere statement that it is part of the invention and reference to a potential method of isolating it. The compound itself is required. See Fiers v. Revel, 25 USPQ2d 1601 at 1606 (CAFC 1993)

and Amgen Inc. v. Chugai Pharmaceutical Co. Ltd., 18 USPQ2II 1016. Therefore, only the isolated nucleic acid set forth in SEQ ID NO: 115, but not the full breadth of the claims meet the written description provision of 35 U.S.C. §112, first paragraph.

Therefore, only the isolated the nucleic acid of SEQ ID NO:421, encoding the polypeptide set forth in SEQ ID NO: 422, but not the full breadth of the claims meet the written description provision of 35 U.S.C. §112, first paragraph.

Claim rejections-35 USC § 112, second paragraph:

The following is a quotation of the second paragraph of 35 U.S.C. 112:

The specification shall conclude with one or more claims particularly pointing out and distinctly claiming the subject matter which the applicant regards as his invention.

6. Claims 119-124 and 128, 132, are rejected under 35 U.S.C. 1 12, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant regards as the invention.

6a. Claims 119-124, 128 and 132 recite ".....the extracellular domain lacking its associated signal sequence....", which renders the claims indefinite because the signal sequence is not generally considered to be part of an extracellular domain, as signal sequences are cleaved from said domains in the process of secretion from the cell. Appropriate correction is required.

Claim Rejections - 35 U.S.C. §102(b):

The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless –

(b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.

7a. Claims 119-138 are rejected under U.S.C. § 102 (b) as being anticipated by Ashkenazi et al (WO200032221; published 08 June 2000).

Ashkenazi et al disclose an isolated nucleic acid that shares 100% homology to the nucleic acid of SEQ ID NO:421 and an isolated polypeptide that shares 100% homology to the polypeptide of SEQ ID NO:422 of the instant application, a vector comprising said nucleic acid, and a host cell comprising said vector. See attached copies of the comparison of SEQ ID NO:421 and SEQ DI NO:422, of the instant invention and the sequence of the reference (SEQUENCE COMPARISON 'A and B", respectively). The nucleic acid disclosed by Ashkenazi et al encodes an isolated polypeptide that lacks its signal sequence. Ashkenazi et al also disclose an isolated nucleic acid that encodes the extracellular domain, (see claims).

Instant claims 119-138 are drawn to an isolated nucleic acid comprising the nucleotide sequence set forth in SEQ ID NO:421, encodes the polypeptide of SEQ ID NO:422, or encoding said polypeptide lacking its signal sequence, or encoding the extracellular domain of the polypeptide of SEQ ID NO:422. Therefore, the Ashkenazi et al reference meets all the limitations recited in claims 119-138, anticipating said claims, in the absence of any evidence to the contrary.

7b. Claims 119-125, 127, 129-137 are rejected under U.S.C. § 102 (b) as being anticipated by Walker et al (WO200029574; published 25 May 2000).

Walker et al disclose an isolated polypeptide that shares 100% homology to the polypeptide of SEQ ID NO:422 and the nucleic acid encoding said polypeptide, a vector comprising said nucleic acid and a host cell comprising said vector. See attached copies of the comparison of SEQ ID NO:422 of the instant invention and the sequence of the reference (SEQUENCE COMPARISON 'C').

Instant claims 119-125 and 130-137 are drawn to an isolated nucleic acid comprising the nucleotide sequence of SEQ ID NO:421, encoding the polypeptide having SEQ ID NO:422, a vector comprising said nucleic acid and a host cell comprising said vector. Therefore, the Walker et al reference meets all the limitations recited in claims 119-125, 130-137, anticipating said claims, in the absence of any evidence to the contrary.

Conclusion:

8. No claim is allowed.

Advisory Information:

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Fozia M Hamud whose telephone number is (571) 272-0884. The examiner can normally be reached on Monday, Thursday-Friday, 6:00 am to 4:00 pm.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Brenda G Brumback can be reached on (571) 272-0961. The fax phone number for the organization where this application or proceeding is assigned is 703-872-9306.

Art Unit: 1647

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see <http://pair-direct.uspto.gov>. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free).

Fozia Hamud
Patent Examiner
Art Unit 1647
15 October 2004



JANET ANDRES
PRIMARY EXAMINER

Sequence Comparison

A 1

Accession	Organism	Sequence
1 CGCTCTGAGTCAGCTGGGGAGTTTCACTTCCCTGGGTCTTCATC	Db	1081 GAGGGGAGAACACATACTCCCRATAATGTCAGGGAGTCAGGAGAAAGAA 1140
QY 61 TGGATTGAAAGTGTGAGAGCAGCATGTTGCCATTGAAACTCTCTGCGTG	QY	1141 CCAGTGAATTCAGGGCCACCTACATGACCATGACCGTGGTGGCTCTGAGG 1200
QY 61 TGGATTGAAAGTGTGAGAGCAGCATGTTGCCATTGAAACTCTCTGCGTG	Db	1141 CCAGTGAATTCAGGGCCACCTACATGACCATGACCGTGGCTCTGAGG 1200
QY 121 TRACTGATATTCTGGGCTGAACTGAACTGATGTTCCCGCTGAGCTAACAGTC	QY	1201 TGGATGGAAACTACTTAAAGTGGGGATGCAAACACAGAA 1260
Db 121 TTRACTGATATTCTGGGCTGAACTGATGTTCCCGCTGAGCTAACAGTC	Db	1201 TGGATGGAAACTACTTAAAGTGGGGATGCAAACACAGAA 1260
QY 181 CAGTGGGGTAGTCACTGGCTGATGGATGTTCCAGAGACAGAGACATGATA	QY	1261 GCCTTGGAGAAGAATGGAGAGTCAGGCTCCCTCATTCAGAGGGGACTCTCTCGTG 1320
QY 241 TCCAAGATAGCTGGACTCTGACACAGAGACCSAACAGGATATGCTATAC	QY	1261 GCCTTGGAGAAGAATGGAGAGTCAGGCTCCCTCATTCAGAGGGGACTCTCTCGTG 1320
QY 241 TCCAAGATAGCTGGACTCTGACACAGAGACCSAACAGGATATGCTATAC	Db	1261 GCCTTGGAGAAGAATGGAGAGTCAGGCTCCCTCATTCAGAGGGGACTCTCTCGTG 1320
QY 301 TATTACCACTCTAGCTGGCTTGGGCTTCAAGAGCTGAGGCTGACAGAATAC	QY	1321 TGTGCTGGCACACTACAGTGAATTCAGACTCCGCTTCCAGCTGCTCCTGT 1380
Db 301 TATTACCACTCTAGCTGGCTTGGGCTTCAAGAGCTGAGGCTGACAGAATAC	Db	1321 TGTGCTGGCACACTACAGTGAATTCAGACTCCGCTTCCAGCTGCTCCTGT 1380
QY 361 GACATTTATGATAGTGGCTCTCTGGCCAAAGTGTGAGGCTGACAGAATAC	QY	1321 TGTGCTGGCACACTACAGTGAATTCAGACTCCGCTTCCAGCTGCTCCTGT 1380
Db 361 GACATTTATGATAGTGGCTCTCTGGCCAAAGTGTGAGGCTGACAGAATAC	Db	1321 TGTGCTGGCACACTACAGTGAATTCAGACTCCGCTTCCAGCTGCTCCTGT 1380
QY 420 361 GACATTTATGATAGTGGCTCTCTGGCCAAAGTGTGAGGCTGACAGAATAC	QY	1381 CTCTTGGAGGACACTTACGGGATGTTGGAGCTGGAGACTGCCTCCAGCTGCTCCTGT 1440
Db 420 361 GACATTTATGATAGTGGCTCTCTGGCCAAAGTGTGAGGCTGACAGAATAC	Db	1381 CTCTTGGAGGACACTTACGGGATGTTGGAGCTGGAGACTGCCTCCAGCTGCTCCTGT 1440
QY 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	1441 ACTCTGGAGGACAGGCTGTGAGGGAGGGAGCTGGAGACTGCCTCCAGCTGCTCCTGT 1500
Db 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	1441 ACTCTGGAGGACAGGCTGTGAGGGAGGGAGCTGGAGACTGCCTCCAGCTGCTCCTGT 1500
QY 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	1501 ACTCTGGCCCTGGAAACAGGTGGAGCTGGGCTCAACCCCGTTGATCAGCC 1560
Db 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	1501 ACTCTGGCCCTGGAAACAGGTGGAGCTGGGCTCAACCCCGTTGATCAGCC 1560
QY 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	1561 CTCCTGGAGGGTTCTAGGAGAGTACTGGAGAGATCAGGATAAACCA 1620
Db 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	1561 CTCCTGGAGGGTTCTAGGAGAGTACTGGAGAGATCAGGATAAACCA 1620
QY 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	1621 CCAAACTAA 1630
Db 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	1621 CCAAACTAA 1630
QY 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	RESULT 2
Db 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	AAT77683
QY 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	ID AAT77683 standard; cDNA; 1630 BP.
Db 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	XX AC AAT77683;
QY 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	XX DT 07-NOV-2000 (first entry)
Db 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	XX DE Human PRO1387 cDNA sequence SEQ ID NO:219.
QY 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	XX KW Human; PRO; promotion; inhibition; angiogenesis; cardiovascularisation; KW diagnosis; trauma; wound; cancer; atherosclerosis; cardiac hypertrophy; KW angiogenic; proliferative; cardiologist; cardiovascular; antiatherosclerotic; KW cytostatic; gene therapy; vaccine; ss.
Db 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	XX OS Homo sapiens.
QY 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	XX PN WO2000032221-A2.
Db 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	XX PD 08-JUN-2000.
QY 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	XX PF 30-NOV-1999; 99WO-0028313.
Db 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	XX PR 01-DEC-1998; 98WO-US025108.
QY 960 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	PR 16-DEC-1998; 98US-0112830P.
Db 960 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	PR 12-JAN-1999; 99US-0115544P.
QY 1020 960 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	PR 08-MAR-1999; 99WO-US005058.
Db 1020 960 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	PR 12-MAR-1999; 99US-0123937P.
QY 1080 1020 960 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	PR 28-APR-1999; 99US-0131445P.
Db 1080 1020 960 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	PR 14-MAY-1999; 99US-013428P.
QY 1080 1020 960 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	PR 03-JUN-1999; 99WO-US01252.
Db 1080 1020 960 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	PR 23-JUN-1999; 99US-0141037P.
QY 1140 1080 1020 960 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	QY	PR 20-JUL-1999; 99US-0144738P.
Db 1140 1080 1020 960 900 840 780 720 660 600 540 480 421 ACCTAACTCTGAAATCGGCCCTCAAGGGAGAGCCAGGGGTGTCACAAGGGGGCTA	Db	PR 26-JUL-1999; 99US-0145688P.

PR 01-SEP-1999; 99WO-US020111.
PR 08-SEP-1999; 99WO-US020594.

Sequence Comparison

Db 421 ACCTATATCTGTGAAATTGGCCCTCAAAAGGGAGGCCAGGGTTCAGGAGCGGTGTTGAGGGGCTGTA 480
 Qy 481 CTGCGATGTCGTCAGAGGCCAAAGAGCTATGTCAGTGTTGGGTATGTGATTCTG 540
 Db 481 CTGCGATGTCGTCAGAGGCCAAAGAGCTATGTCAGTGTTGGGTATGTGATTCTG 540
 Db 541 ATGGGTTGTTTCCAGAGCAGAGCTGAAACCTGACCAAGTTAGATGGTATT 600
 Qy 541 ATGGGTTGTTTCCAGAGCAGAGCTGAAACCTGACCAAGTTAGATGGTATT 600
 Db 601 TCAGSAGGGGCGGCGAAGAGGAGATGAAAGCTGCTATGGGATGTTAGGATCT 660
 Qy 601 TCAGSAGGGGCGGCGAAGAGGAGATGAAAGCTGCTATGGGATGTTAGGATCT 660
 Db 661 GTGGGACTCCCGAGCTGGGACATTCAGAGTGTGACCTGGTGGGGATT 720
 Qy 661 GTGGGACTCCCGAGCTGGGACATTCAGAGTGTGACCTGGTGGGGATT 720
 Db 661 GTGGGACTCCCGAGCTGGGACATTCAGAGTGTGACCTGGTGGGGATT 720
 Qy 721 TTCCGAATGCGGTTCCATCATGCTTAAGGGTGTGGACTCATGGAAACTAC 780
 Db 721 TTCCGAATGCGGTTCCATCATGCTTAAGGGTGTGGACTCATGGAAACTAC 780
 Qy 781 ACCTGAGTATCCACCTAGGAACCTGGTCAAGAAACATTGCTGCGATGAGC 840
 Db 781 ACCTGAGTATCCACCTAGGAACCTGGTCAAGAAACATTGCTGCGATGAGC 840
 Qy 841 CGGGAGAGCTCGAACATGGTACCCGGAGCCCTGGCTTCAGAAACCTGGCTTGGGAGCT 900
 Db 901 AACATCGTGGTGCATATGGTGCATATGGTCTGGCCCAAACCTGTCCTGGCTTGGGAGCT 900
 Db 901 AACATCGTGGTGCATATGGTGCATATGGTCTGGCCCAAACCTGTCCTGGCTTGGGAGCT 900
 Qy 961 ATATTCATCTGGAGAGACCTGGAAATAAGAGTCAGTGAATCTACGTCTGG 1026
 Db 961 ATATTCATCTGGAGAGACCTGGAAATAAGAGTCAGTGAATCTACGTCTGG 1026
 Qy 1021 AAGACAGAGAGAGATATCGAGATAAAGAAAACCTGGCTTGTAGATG 1084
 Db 1021 AAGACAGAGAGAGATATCGAGATAAAGAAAACCTGGCTTGTAGATG 1084
 Qy 1081 GAGGGAGAACACTTACTCCATAATGTCGGGGTGTGAGGAAGAGAA 1140
 Db 1081 GAGGGAGAACACTTACTCCATAATGTCGGGGTGTGAGGAAGAGAA 1140
 Qy 1141 CCAGGGAATTCAGGGCCACTATGACATGGACCCGTTGGCTTCTGAGG 1200
 Db 1141 CCAGGGAATTCAGGGCCACTATGACATGGACCCGTTGGCTTCTGAGG 1200
 Qy 1201 TCGATCTGGACACACTCACTTGAAALAAAGTCAGGTTGGGATGCAAAACAGCAA 1260
 Db 1201 TCGATCTGGACACACTCACTTGAAALAAAGTCAGGTTGGGATGCAAAACAGCAA 1260
 Qy 1261 GCCTTGTGAGAGATGGAGTCCTCATCTGGAGACTCTCTCTGGTG 1320
 Db 1261 GCCTTGTGAGAGATGGAGTCCTCATCTGGAGACTCTCTCTGGTG 1320
 Qy 1321 TGTGTTCTGGCCACTCTCCAGTGTGAACTGAGCTGGGCTTCCAGCTCTCTGGTG 1380
 Db 1321 TGTGTTCTGGCCACTCTCCAGTGTGAACTGAGCTGGGCTTCCAGCTCTCTGGTG 1380
 Qy 1381 CTGATGTTGTCATACACTGAGATGGAGATGGGATCTGAGCTGGGAGCTGGG 1440
 Db 1381 CTGATGTTGTCATACACTGAGATGGAGATGGGATCTGAGCTGGGAGCTGGG 1440
 Qy 1441 AGCTCTGGAGAACGSGCTGCTGAGGGAGGGAGCATGACTCTGGCTTCTGGTG 1500
 Db 1441 AGCTCTGGAGAACGSGCTGCTGAGGGAGGGAGCATGACTCTGGCTTCTGGTG 1500
 Qy 1501 ACATGGCCCTGGGACCCAGGCTGAGCTGGCTCAACCCCCGGTGTGATGAC 1560

Sequence comparison

B

PR	13-SEP-1999;	99WO-US020944.	ID	AU1231 standard; protein; 394 AA.
PR	15-SEP-1999;	99WO-US021547.	XX	
PR	05-OCT-1999;	99WO-US023059.	AC	AU1231;
PR	29-OCT-1999;	99US-0162506P.	XX	
XX			DT	24-OCT-2001 (first entry)
PA	(GETH) GENENTECH INC.		XX	Human PRO1387 polypeptide sequence.
XX			DE	
PI	Aebkenazi AJ, Baker JP, Ferrara N, Gerber H, Hillan KJ, Klein RD, Kuo SS, Paoni NF,	KW	Human secretory and transmembrane; PRO; mammalian; cancer; lung; breast;	
PI	Goddard A, Godowski PJ, Gurney AL, Klein RD, Kuo SS, Paoni NF,	KW	prostata; cervical; tumour necrosis factor-alpha; tnf-alpha; cartilage;	
PI	Smith V, Watanabe CK, Williams PM, Wood WI,	KW	ear; proliferation; glucose; free fatty acid; skeletal muscle; adipocyte;	
XX		KW	A-peptide; factor VIIa; gene therapy.	
DR	WPI: 2000-412154/35.	OS	Homo sapiens.	
DR	N-PSDB; AAS177683.	XX		
XX		XX		
PT	Nucleic acids encoding PRO polypeptides useful for preventing, diagnosing	PN	W0200140466-A2.	
PT	and treating disorders in cardiovascular, endothelial or angiogenic	XX		
XX	disorders in mammals.	PD	07-JUN-2001.	
PS		XX		
Claim 72; Fig 92; 315pp; English.		01-DEC-2000; 2000WO-US032678.		
XX				
CC	The present invention describes nucleic acids encoding PRO polypeptides	PR	01-DEC-1999; 99WO-US038301.	
CC	useful for preventing, diagnosing and treating disorders in	PR	01-DEC-1999; 99WO-US038634.	
CC	cardiovascular, endothelial or angiogenic disorder in mammals by	PR	02-DEC-1999; 99WO-US028551.	
CC	modulating cell proliferation, angiogenesis and cardiovascularisation,	PR	02-DEC-1999; 99WO-US038565.	
CC	and for identifying agonists and antagonists of these processes. The	PR	09-DEC-1999; 99US-0170262P.	
CC	nucleic acids and the proteins they encode may be used in the prevention,	PR	16-DEC-1999; 99WO-US030095.	
CC	treatment and diagnosis of diseases associated with inappropriate PRO	PR	20-DEC-1999; 99WO-US030911.	
CC	expression such as cardiovascular, endothelial or angiogenic disorders in	PR	20-DEC-1999; 99WO-US030999.	
CC	mammals (e.g., atherosclerosis, cancers, and cardiac hypertrophy). For	PR	30-DEC-1999; 99WO-US031243.	
CC	example, the nucleic acids (NCs) and vector containing them and the PRO	PR	30-DEC-1999; 99WO-US031274.	
CC	polypeptide may be used to treat disorders associated with decreased PRO	PR	05-JAN-2000; 2000WO-US000219.	
CC	expression. AAS177510 to AAS17721 and AAS24388 represent decreased	PR	06-JAN-2000; 2000WO-US000277.	
CC	nucleotide and protein sequences used in the exemplification of the	PR	06-JAN-2000; 2000WO-US000376.	
CC	present invention.	PR	11-FEB-2000; 2000WO-US000365.	
XX		PR	18-FEB-2000; 2000WO-US004341.	
SQ	Sequence 394 AA;	PR	22-FEB-2000; 2000WO-US004342.	
Query Match	100.0%; Score 2667; DB 3; Length 394;	PR	24-FEB-2000; 2000WO-US004114.	
Best Local Similarity	100.0%; Fred. No. 5.1e-188;	PR	24-FEB-2000; 2000WO-US005004.	
Matches	394; Conservative 0; Mismatches 0; Indels 0; Gaps 0;	PR	01-MAR-2000; 2000WO-US005601.	
QY	1 MFCQLKLILPLVLDYSDIGLNDINVSPELTWVGDALMGCVFQSTEDKCFIKDWTS 60	PR	03-MAR-2000; 2000WO-US005841.	
Db	1 MPCQLKLILPLVLDYSDIGLNDINVSPELTWVGDALMGCVFQSTEDKCFIKDWTS 60	PR	10-MAR-2000; 2000WO-US006319.	
QY	61 PGERHAKDETYLYYYNSLSPVIGRQNRYFLGDLICNDSLILQDQBAOGTYCIEIR, 120	PR	15-MAR-2000; 2000WO-US006884.	
Db	61 PGERHAKDETYLYYYNSLSPVIGRQNRYFLGDLICNDSLILQDQBAOGTYCIEIR, 120	PR	20-MAR-2000; 2000WO-US007377.	
QY	121 KGEQVKPKAVVLLVLPSPERKEIYVHGGLIGNCVQSTETKVKVENVIFSGRAKEE 180	PR	21-MAR-2000; 2000WO-US007532.	
Db	121 KGEQVKPKAVVLLVLPSPERKEIYVHGGLIGNCVQSTETKVKVENVIFSGRAKEE 180	PR	30-MAR-2000; 2000WO-US008439.	
QY	121 KGEQVKPKAVVLLVLPSPERKEIYVHGGLIGNCVQSTETKVKVENVIFSGRAKEE 180	PR	17-MAY-2000; 2000WO-US010705.	
Db	121 KGEQVKPKAVVLLVLPSPERKEIYVHGGLIGNCVQSTETKVKVENVIFSGRAKEE 180	PR	22-MAY-2000; 2000WO-US014042.	
QY	181 IFRYTHKRLMSVSYQSQGHFQNRVNLGDFRNDSGIMLQGVRSQDGNYTCIHLGN 240	PR	30-MAY-2000; 2000WO-US014941.	
Db	181 IFRYTHKRLMSVSYQSQGHFQNRVNLGDFRNDSGIMLQGVRSQDGNYTCIHLGN 240	PR	05-JUN-2000; 2000WO-US015264.	
QY	241 LVFKTIVHVSPEPRTVTPALRPVLTGSQVLTIVGVCATILLVPLILVVKTC 300	PR	05-JUN-2000; 2000WO-US020832P.	
Db	241 LVFKTIVHVSPEPRTVTPALRPVLTGSQVLTIVGVCATILLVPLILVVKTC 300	PR	20-JUL-2000; 2000WO-US020710.	
QY	301 GNKSYSNTVLUKQMKKPKPEIKEPKCPREREGEKGHISPIVREVIEWEPEKSEKT 360	PR	13-AUG-2000; 2000WO-US022031.	
Db	301 GNKSYSNTVLUKQMKKPKPEIKEPKCPREREGEKGHISPIVREVIEWEPEKSEKT 360	PR	23-AUG-2000; 2000WO-US023522.	
QY	361 YMTMHPWVMSLRDRNNSLERKGGGMKTTQCAF 394	PR	24-AUG-2000; 2000WO-US023128.	
Db	361 YMTMHPWVMSLRDRNNSLERKGGGMKTTQCAF 394	PR	05-NOV-2000; 2000WO-US030952.	
RESULT 5	AAU12431	PR	10-NOV-2000; 2000WO-US030873.	
AAU12431		PA	(GETH) GENENTECH INC.	
		XX		
		PI	Baker JP, Beresini M, Deforge L, Destroyers L, Filvaroff E, Gao W;	
		PI	Gerritsen ME, Goddard A, Godowski PJ, Gurney AL, Sherwood S;	
		PI	Smith V, Stewart TA, Tumas D, Watanabe CK, Wood WI, Zhang Z;	
		XX		
		DR	WPI; 2001-408281/43.	
		XX		
		PT	Isolated, secretory and transmembrane PRO polypeptide used to detect other PRO polypeptides, link bioactive molecules to cells expressing PRO	

Sequence Comparison

Db 61 PGEHAKDSEVLYTTSNLSPIGRFQNRVHLMDLICNDGSLIQQDVOBADOGTYICEIRL 120
QY 121 KGSQVFKKAVVHLWLPBPKELMWHGGIQLOMGCVFQSTEVKETVKVWIFSGRRAKEE 180

Db
QY
121 KGBESQVKKAVLHTRPESKELMNVHGGLIOMGCVFQSTEVTHVKEVNPGRKEE 180
181 IFRYHKLMNSVEYSQSMWHFONVNLGDIFRNDSDIMLOGVREDDGTYCTCILHGN 240

Db	<p>181 IYFRYTHKLRSVYESQSWMHFQNRVNVLGDDIFRNDISIMLOGVRESDGGNYTCIHLGN 240</p> <p>QY </p> <p>241 LVPKKTIVLWSPERRTLTVPAILRPLVIGGMQNLVINGIVCATILLPVLILVKTC 300</p> <p>Db 241 LVPKKTIVLWSPERRTLTVPAILRPLVIGGMQNLVINGIVCATILLPVLILVKTC 300</p> <p>QY </p> <p>301 GNKSSVNSTLVLKNTRKTPBEIKRPCHFRCGEGKHYSPIVREVIEEPSEKSEAT 360</p> <p>Db 301 GNKSSVNSTLVLKNTRKTPBEIKRPCHFRCGEGKHYSPIVREVIEEPSEKSEAT 360</p> <p>QY </p> <p>361 YNTMHPWPSLRSDRNLSLEKKGGMPKTQAF 394</p> <p>Db 361 YNTMHPWPSLRSDRNLSLEKKGGMPKTQAF 394</p>
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Sequence compare
C)
RESULT 3
RAY9452
ID RAY9452 standard; protein; 394 AA.
XX
AC RAY9452;
XX
DT 11-SEP-2000 (first entry)
XX
DE Human inflammation associated protein #11.
XX
KW Inflammation; rheumatoid arthritis; Crohn's disease; asthma;
KW multiple sclerosis; allergy; AIDS; diabetes mellitus; antiinflammatory;
GW gene therapy; human

Query Match CC #11, derived from Incyte Clone 3507924
 Pct. ID Similarity 100.0% Score 206
 Pct. Gln Similarity 100.0% Score 206
 XX SQ Sequence 394 AA;

Matches	Similarity	Length	Start	End	Indels	Gaps
QY	Conservative	0	0	0	0	0
Db						
1	MPCPLKLILPVLVDYLDSIGLNDLNVSPELTWVGDALNGCVFQSTEDKCIFKIDWLS	60				
1	MPCPLKLILPVLVDYLDSIGLNDLNVSPELTWVGDALNGCVFQSTEDKCIFKIDWLS	60				
QY						
61	PGEHAKEDEVLYYNSNVSPIRFQNRVHMGDILNDGSILQVOADQGYCIEIRL	120				
61	PGEHAKEDEVLYYNSNVSPIRFQNRVHMGDILNDGSILQVOADQGYCIEIRL	120				
Db						
121	KGESEQVKAWIHLVPEPKELMVRGLQMGCVFQSTEVKHTVKENIFSGRKEE	180				
121	KGESEQVKAWIHLVPEPKELMVRGLQMGCVFQSTEVKHTVKENIFSGRKEE	180				
Db						
181	IYFRYHKLMSYESQSQWGFQNRYVLDIPRNDSIMQGVRSBDSGANTCISHLGN	240				
181	IYFRYHKLMSYESQSQWGFQNRYVLDIPRNDSIMQGVRSBDSGANTCISHLGN	240				
Db						
241	IYFKKTIVLAVSBEPRNTPAALRIVLGENQNTIVGIVCATILLPVLLIVKTC	300				
241	IYFKKTIVLAVSBEPRNTPAALRIVLGENQNTIVGIVCATILLPVLLIVKTC	300				
Db						
301	GINKSSWNTLVKOTKINPTEIKEPKHFERGERGERKHYTIVRIVEEPPSKSEAT	360				
301	GINKSSWNTLVKOTKINPTEIKEPKHFERGERGERKHYTIVRIVEEPPSKSEAT	360				
QY						
361	YMTMPFWVSLRSDRNNSLEKKGGGPKTOAF	394				
361	YMTMPFWVSLRSDRNNSLEKKGGGPKTOAF	394				

Sequence Comparison

DE	XX	Human PRO1387 protein sequence SEQ ID NO:220.
KW	XX	Human; PRO; promotion; inhibition; angiogenesis; cardiovascularisation;
KW	XX	diagnosis; trauma; wound; cancer; atherosclerosis; cardiac hypertrophy;
KW	XX	angiogenic; proliferative; cardiot; cardiovascular; antiatherosclerotic;
KW	XX	cytostatic; gene therapy; vaccine.
OS	XX	Homo sapiens.
PN	XX	WO20032221-A2.
PR	XX	08-JUN-2000.
PR	XX	08-NOV-1999;
PP	XX	99W0-US028313.
PR	01-DEC-1998;	99W0-US025108.
PR	16-DEC-1998;	98US-0112850P.
PR	16-JUN-1999;	99US-0115554P.
PR	08-MAR-1999;	99W0-US05028.
PR	12-MAR-1999;	99US-0123957P.
PR	28-APR-1999;	99US-0131445P.
PR	14-MAY-1999;	99US-0134287P.
PR	02-JUN-1999;	99W0-US01225Z.
PR	23-JUN-1999;	99US-0141037P.
PR	20-JUL-1999;	99US-0144758P.
PR	26-JUL-1999;	99US-0145698P.
PR	01-SEP-1999;	99W0-US020111.
PR	08-SEP-1999;	99W0-US020594.